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(71) Applicant (for all designated States except US): **HETERO DRUGS LIMITED** [IN/IN]; Hetero House, 8-3-166/7/1, Erragadda, Hyderabad, Andhrapradesh, Hyderabad 500018 (IN).

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(72) Inventors; and

(75) Inventors/Applicants (for US only): **PARTHASARADHI REDDY, bandi** [IN/IN]; Hetero House, 8-3-166/7/1, Erragadda, Hyderabad, Andhrapradesh, Hyderabad 500018 (IN). **RATHNAKAR REDDY, kura** [IN/IN]; Hetero Drugs Limited (R & D), Plot No. B-80 & 81, A.P.I.E., Balanagar, Hyderabad 500018 (IN). **RAJI REDDY, rapolu** [IN/IN]; Hetero Drugs Limited (R & D), Plot No. B-80 & 81, A.P.I.E., Balanagar, Hyderabad 500018 (IN). **MURALIDHARA REDDY, dasari** [IN/IN]; Hetero Drugs Limited (R & D), Plot No. B-80 & 81, A.P.I.E., Balanagar, Hyderabad 500018 (IN). **SRINIVAS REDDY, itiyala** [IN/IN]; Hetero House, 8-3-166/7/1, Erragadda, Hyderabad, Andhrapradesh, Hyderabad 500 018 (IN).

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(54) Title: A NOVEL PROCESS FOR PREPARATION OF NEBIVOLOL INTERMEDIATES

(57) Abstract: The present invention relates to a process for separation of desired diastereomeric pair from a mixture of diastereomeric pairs thereby obtaining nebivolol intermediates. Thus, the mixture of (+)-[IS*(R*)]-6-fluoro-3,4-dihydro- α -[[(phenylmethyl)amino]methyl]-2H-l-benzopyran-2-methanol, (+)-[IS*(S*)]-6-fluoro-3,4-dihydro-2-oxiranyl-2H-l-benzopyran and ethanol is heated to reflux temperature and stirred for 8 hours at the same temperature to obtain (±)-[2R*[IS*,5S*(S*)]]+[2R*[IS*,5R*(R*)]]- α,α' -[phenylmethyliminobis(methylene)]bis[6-fluoro-3,4-dihydro-2H-l-benzopyran-2-methanol]. Then the reaction mass is cooled to 10°C, the pH is adjusted to 2 with HCl gas and stirred for 45 minutes at 25°C to 30°C. Then the separated solid is filtered and dried to give (+)-[2R*[IS*,5S*(S*)]]- α,α' -[phenylmethyliminobis(methylene)]bis[6-fluoro-3,4-dihydro-2H-l-benzopyran-2-methanol] hydrochloride salt, which can be converted into nebivolol.

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